

Jacques Monod (1910-1976) and his publications in the "Annales de l'Institut Pasteur".

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17 Abstract

Between 1942 and 1956, Jacques Monod, Nobel Prize winner in Physiology or Medicine, contributed a number of papers to the *Annales de l' Institut Pasteur*, the ancestor of the journal "Research in Microbiology". Circumstances that led him to publish in the "*Annales*" are recalled here.

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Keywords: Jacques Monod; Nobel Prize; Annales de l'Institut Pasteur; Diauxic growth;
Enzymatic adaptation

1. Introduction

2010 is the centenary anniversary of the birth of Jacques Monod, who together with François Jacob and André Lwoff received the Nobel Prize in 1965 in recognition of their outstanding contribution to molecular biology. There exists abundant literature concerning the various stages of his life and his scientific achievements (1, 5, 6, 8-10), but the early stages of his scientific career are generally not emphasized. It so happens that, between 1942 and 1956, Jacques Monod published 16 articles in the "Annales de l'Institut Pasteur" (3). His first publications in the "Annales" were not symbolic of his work, but they represent a decisive turning point in his intellectual pursuit which was to lead to the most prestigious recognition in the international scientific world.

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39 Monod, who obtained a science degree (licence) in Paris in 1931, was granted a 40 scholarship so as to learn microbiological techniques at Strasbourg University (France), in the 41 laboratory of Edouard Chatton, a specialist in protozoa. The following year he returned to 42 Paris on a Commercy fellowship to join the "Laboratoire d'évolution des êtres organisés" 43 (Laboratory of evolution of living organisms), directed by his former professor at the 44 Sorbonne, Maurice Caullery, and continued working with ciliates. Then, in 1934, he became 45 an assistant in the laboratory of Zoology (Faculty of Sciences, Paris) directed by Charles 46 Perez. It was to be his first paid permanent position. But the job did not suit him well-- he 47 worked in a small dilapidated room, with poor facilities and little exchange with his 48 colleagues.

49 To get away from his dreary laboratory he joined Paul-Emile Victor in a scientific 50 expedition to Greenland on the "Pourquoi pas?" He then had the opportunity of learning 51 genetics. He obtained a Rockefeller grant, in 1936 to work on eye pigmentation genes of 52 Drosophila in the laboratory of Thomas H. Morgan at the California Institute of Technology 53 in Pasadena (Fig. 1). He discovered another way to perform research founded on collaborative 54 work and critical and fruitful discussions. Back in France, he carried on research for a while at 55 the Institute of Physicochemical Biology, where he continued to work on the model 56 developed in USA.

57 Having kept his position of assistant at the Sorbonne, Jacques Monod came back to 58 work in the laboratory of Charles Perez in 1937, where he began his thesis on bacterial 59 growth using Escherichia coli as a model system. This choice resulted form the influence of 60 two men. As a student in zoology, it was during a training course in the Laboratory of Marine 61 Biology in Roscoff (France) in 1929 that Jacques Monod met Georges Teissier for the first 62 time. The latter was a specialist in biometrics, and oriented Jacques Monod's research 63 towards the study of the kinetics of the growth of bacteria. It was also in Roscoff that Monod 64 met André Lwoff. Lwoff, who was head of the Service of Microbial Physiology at the Institut 65 Pasteur, was well known for his work on growth factors. He convinced Monod to give up the 66 ciliate model for bacteria and provided him with a culture of *E. coli*.

67 Thus, when he decided to begin his thesis, Monod had already acquired wide 68 experience working in many laboratories. This propensity to move from one laboratory to 69 another without settling in reflects the state of mind of Monod at the time, for he was also tempted by a career in music, for which he was extremely gifted. Jacques Monod took histime, but finally chose to pursue his career in biological research.

72 During the Second World War, Monod joined the French Resistance. His laboratory 73 harbored a secretarial office for printing up anti-Nazi pamphlets. Although his commitment 74 considerably slowed down his scientific activities, he managed to finalize his thesis dissertation, which he defended in 1941 under the title "Recherches sur la croissance des 75 cultures bactériennes". In his memoir published in 1942 (7) he reported on an unknown 76 77 phenomenon which he termed "diauxic growth" to account for the fact that when bacteria are 78 placed in a medium containing both glucose and lactose, growth is continuous until depletion 79 of glucose; then, after a pause, growth resumes with the use of lactose. André Lwoff then 80 pointed out to him that this phenomenon was certainly a particular example "of enzymatic 81 adaptation". This work also led to a publication in the "Annales de l'Institut Pasteur" in 1942 82 (6). Since he was being spied upon by the Gestapo due to his active role in the French 83 Resistance, he left the Sorbonne and took refuge at the Institut Pasteur in André Lwoff's laboratory to study enzymatic adaptation. After the liberation, he engaged in the 1st French 84 85 Army of General de Lattre de Tassigny. He officially joined the Institut Pasteur as Head of a 86 Laboratory in 1945, where he worked until his death in 1976.

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88 **3.** Publication in the "Annales"

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90 Under what circumstances did Monod choose to publish his work in the "Annales de 91 l'Institut Pasteur"? The "Association des Microbiologistes de Langue Française" (AMLF), 92 which later became the French Society for Microbiology, was created in 1938 (2). Scientific 93 reports presented in the format of research papers (i.e. including methods, tables and figures) 94 were usually published in the "Annales". During the War, the AMLF had difficulty in 95 organizing congresses, and the Institut Pasteur held monthly meetings on its premises. André 96 Lwoff, who was an AMLF member, encouraged Monod to present his data. Until then, 97 Jacques Monod's works had always been presented during formal sessions of the French 98 Academy of Sciences. He thus discovered a more casual atmosphere and an audience which 99 greatly appreciated the originality of his discoveries. This is how 10 of his contributions were 100 published in the "Annales de l'Institut Pasteur", including a paper with Elie Wollman (Table 101 1). The initial objective of the "Annales" was to publish research work performed in the Institut Pasteur laboratories (both on the mainland and overseas) (3). It was thus within thisframework that Jacques Monod published 6 other articles (Table 1).

104 While Monod regularly contributed papers to the "Annales" between 1942 and 1947, 105 only three papers were published in the journal thereafter (Table 1). The explanation for this 106 is simple. French continued to be the main language of the "Annales" and Monod's work, 107 highly specialized in genetics and molecular biology, addressed an international readership. 108 He thus published in Anglo-Saxon journals. Two of those three papers published during the 109 1950s included results from projects funded by foreign institutions (Table 1: papers 14 and 110 16). Since the second one was also sponsored by a French institution, it was important to 111 publish the results in a French journal. In the third article, Monod returned to the source of 112 his works: bacterial growth. In that paper, Monod set out the theoretical and experimental 113 bases of a method of continuous culture (Table 1: paper 15). It is understandable that this 114 particular work was published in the Annales, since Monod had been teaching microbiology 115 courses since 1945. The experimental possibilities offered by this new technique were to be 116 widely used thereafter in French and foreign laboratories.

After his death, several of his papers that had been published in the "*Annales*" were reproduced in a volume of selected papers edited by André Lwoff and Agnes Ullmann (4, Table 1), bearing witness to the importance of his contributions. He himself cited three of the papers published in the "*Annales*" (Table 1) in his Nobel Lecture (8).

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122 **4.** Conclusion

Jacques Monod's papers published in the "*Annales de l'Institut Pasteur* " came just after his work on bacterial growth and were a continuation of his thesis. Since they were at the origin of the scientific work which led him to the Nobel Prize, they constitute pioneering research and are important in understanding the future orientation of his research, which was determinant for the continuation of his career: in particular, the discovery of the transcriptional regulation system, in which he analyzed control of genetic expression in the *lac* operon, in *Escherichia coli* and the theory of allosteric interactions (8).

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131 Acknowledgements

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| 162 163 164 165 | Legend Fig. 1: Jacques Monod (1910-1976) in Pasadena, California in 1936. | |
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168 Table 1 Work of Jacques Monod published in the "Annales de l'Institut Pasteur"

| | Туре | Authors | Title | Year, volume, pages |
|-----|--------------------------------------------------|----------------------------------------|-------------------------------------------------------------------------------------------------------------------------------------------|------------------------|
| 1. | AMLF report 7 May 1942 | Monod, J. | Influence de l'amide de l'acide nicotinique, de l'aneurine et de l'acide ascorbique sur la croissance d' <i>Escherichia coli</i> | (1942) 68, 435- 438 |
| 2. | AMLF report 5 March 1942 | Monod, J. | Sur le phénomène de lyse lié à l'inanition carbonée | (1942) 68, 445- 451 |
| 3. | ALMF report 2 July 1942 ^a | Monod, J. | Diauxie et respiration au cours de la croissance des cultures de <i>B. coli</i> | (1942) 68, 548- 450 |
| 4. | AMLF report 4 February 1943 | Monod, J. | Inflence de la concentration des substrats sur la rapidité d'adaptation chez le <i>B. coli</i> | (1943) 69, 179- 181 |
| 5. | AMLF report 1 July 1943 | Monod, J. | Sur la non-additivité d'action de certains enzymes bactériens | (1944) 70, 57-59 |
| 6. | AMLF report 1 July 1943 | Monod, J. | Remarque sur le problème de la spécificité des enzymes bactériens | (1944) 70, 60-61 |
| 7. | AMLF report 1 December 1943 ^{a,b} | Monod, J. | Inhibition de l'adaptation enzymatique chez <i>B. coli</i> en présence de 2-4 dinitrophénol | (1944) 70, 381- 384 |
| 8. | AMLF report 2 December 1943 ^a | Monod, J. | Sur la nature du phénomène de diauxie | (1945) 71, 37-38 |
| 9. | AMLF report 8 November 1945 | Morel. M., Monod, J. | Sur l'utilisation du saccharose par Proteus vulgaris | (1946) 72, 647- 656 |
| 10. | Research paper ^{a, b} | Monod, J., Audureau, A. | Mutation et adaptation enzymatique chez <i>Escherichia</i> <i>coli</i> -mutabile | (1946) 72, 868- 878 |
| 11. | Research paper ^a | Monod, J. | Sur une mutation spontanée affectant le pouvoir de synthèse de la méthoinine chez une bactérie coliforme | (1946) 72, 879- 890 |
| 12. | Research paper | Lwoff, A., Monod J. | Essai d'analys edu rôle de l'anhydride carbonique dans la croissance microbienne | (1947) 73, 323- 347 |
| 13. | AMLF report 3 April 1947 | Monod, J., Wollman, E. ^a | L'inhibition de la croissance et de l'adaptation enzymatique chez les bactéries infectées par le bacteriophage | (1947) 73, 937- 956 |
| 14. | Research paper ^{a, c} | Monod, J., Torriani, A. | De l'amylomaltase d' <i>Escherichia</i> coli | (1950) 78, 65-77 |
| 15. | Research paper ^a | Monod, J. | La technique de culture continue: théorie et applications | (1950) 79, 390- 410 |
| 16. | Research paper ^{a, b,d} | Rickenberg, H., Cohen | La galactoside-perméase d'Escherichia coli | (1956) 91, 829- 857 |

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- AMLF: Association des Microbiologistes de Langue Française

- ^a Work reproduced in the selected papers (4)
 ^b Work cited in the Nobel lecture (8)
 ^c Work funded by the National Institute of Health
- ^d Work funded by The Rockefeller Foundation, Jane Coffin Child's Memorial Fund
- and the "Commissariat à l'Energie Atomique"

